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Attorney Docket No. P67772US1
Application No. 10/509,950**Amendments to the claims:**

This listing of claims replaces all prior versions, and listings, of claims in the application.

Listing of claims:

Claims 1-10 (cancelled).

11 (withdrawn): An antibody specifically immunoreactive with an immunogen, wherein said immunogen is a protein molecule shown in SEQ ID NO. 1, or a fragment, or derivative, or variant thereof.

12 (withdrawn): Use of an antibody of claim 11, for detecting the pathological state of a cell in a sample from a subject, comprising immunocytochemical staining of said cell with said antibody, wherein an altered degree of staining, or an altered staining pattern in said cell compared to a cell representing a known health status indicates a pathological state of said cell.

13 (currently amended): A method of diagnosing ~~or prognosticating~~ Alzheimer's disease in a subject, ~~or determining whether a subject is at increased risk of developing said disease,~~ comprising:

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- determining a ratio level ~~and/or an activity~~ of
- (i) a transcription product of the gene coding for polypeptide hTARPP (SEQ ID NO: 1), ~~and/or~~
- (ii) a translation product of the gene coding for Htarpp (SEQ ID NO: 1),
- in a brain tissue sample of temporal cortex, frontal cortex, and/or hippocampus obtained from said subject and
- comparing said ratio level ~~and/or said activity~~ to a reference value representing a known disease or health status, thereby diagnosing ~~or prognosticating~~ Alzheimer's disease in said subject, ~~or determining whether said subject is at increased risk of developing Alzheimer's disease.~~

Claims 14-17 (cancelled).

- 18 (currently amended): The method according to claim 13 wherein said reference value representing a known health status is that of a ratio level ~~and/or an activity~~ of
- (i) a transcription product of the gene coding for polypeptide hTARPP (SEQ ID NO: 1) ~~and/or~~
- (ii) a translation product of the gene coding for hTARPP (SEQ ID NO: 1),
- in a brain tissue sample of temporal cortex, frontal cortex, and/or hippocampus sample obtained from a subject not suffering from Alzheimer's disease.

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19 (currently amended): The method according to claim 13 wherein an alteration in the ratio level and/or activity of a transcription product of the gene coding for polypeptide hTARPP (SEQ ID NO: 1) and/or a translation product of the gene coding for hTARPP (SEQ ID NO: 1) in a brain tissue sample of temporal cortex, frontal cortex, and/or hippocampus cell, or tissue, or fluid obtained from said subject relative to a reference value representing a known health status indicates a diagnosis, or prognosis, or increased risk of Alzheimer's disease in said subject.

20 (currently amended): A kit for diagnosing or prognosticating Alzheimer's disease in a subject, or determining the propensity or predisposition of a subject to develop Alzheimer's disease, said kit comprising:

(a) at least one reagent ~~which is~~ selected from the group consisting of

—(i) reagents that selectively detect a transcription product of the gene coding for polypeptide hTARPP (SEQ ID NO: 1), and

—(ii) ~~reagents that selectively detect a translation product of the gene coding for hTARPP (SEQ ID NO: 1), and~~

(b) an instruction for diagnosing, or prognosticating Alzheimer's disease or ~~determining the propensity or predisposition of a subject to develop Alzheimer's disease by~~

— detecting a ratio level, or an activity, or both said level and said activity, of said transcription product and/or said translation product of the gene coding for polypeptide

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hTARPP (SEQ ID NO: 1), in a brain tissue sample of temporal cortex, frontal cortex, and/or hippocampus obtained from said subject[;] and

- ~~diagnosing or prognosticating Alzheimer's disease or determining the propensity or predisposition of said subject to develop Alzheimer's disease, wherein a varied (i) when a ratio level, or activity, or both said level and said activity, of said transcription product is varied and/or said translation product compared to a reference value representing a known health status[,]~~ or wherein a (ii) when a ratio level, or activity, or both said level and said activity, of said transcription product and/or said translation product is similar or equal to a reference value representing a known disease status indicates a diagnosis or prognosis of Alzheimer's disease or an increased propensity or predisposition of developing Alzheimer's disease.

- 21 (withdrawn): A method of treating or preventing a neurodegenerative disease, in particular Alzheimer's disease, in a subject comprising administering to said subject in a therapeutically or prophylactically effective amount an agent or agents which directly or indirectly affect an activity and/or a level of (i) a gene coding for hTARPP, and/or (ii) a transcription product of the gene coding for hTARPP, and/or (iii) a translation product of the gene coding for hTARPP, and/or (iv) a fragment, or derivative, or variant of (i) to (iii).

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- 22 (withdrawn): A modulator of an activity and/or of a level of at least one substance which is selected from the group consisting of (i) a gene coding for hTARPP, and/or (ii) a transcription product of the gene coding for hTARPP, and/or (iii) a translation product of the gene coding for hTARPP, and/or (iv) a fragment, or derivative, or variant of (i) to (iii).
- 23 (withdrawn): Use of a modulator of an activity and/or of a level of at least one substance which is selected from the group consisting of (i) a gene coding for hTARPP, and/or (ii) a transcription product of the gene coding for hTARPP, and/or (iii) a translation product of the gene coding for hTARPP, and/or (iv) a fragment, or derivative, or variant of (i) to (iii) for a preparation of a medicament for treating or preventing a neurodegenerative disease, in particular Alzheimer's disease.
- 24 (withdrawn): A recombinant, non-human animal comprising a non-native gene sequence coding for hTARPP or a fragment, or a derivative, or a variant thereof, said animal being obtainable by:
- (i) providing a gene targeting construct comprising said gene sequence and a selectable marker sequence, and
 - (ii) introducing said targeting construct into a stem cell of a non-human animal, and
 - (iii) introducing said non-human animal stem cell into a non-human embryo, and
 - (iv) transplanting said embryo into a pseudopregnant non-human animal, and
 - (v) allowing said embryo to develop to term, and

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- (vi) identifying a genetically altered non-human animal whose genome comprises a modification of said gene sequence in both alleles, and
 - (vii) breeding the genetically altered non-human animal of step (vi) to obtain a genetically altered non-human animal whose genome comprises a modification of said endogenous gene, wherein said disruption results in said non-human animal exhibiting a predisposition to developing symptoms of a neurodegenerative disease or related diseases or disorders.
- 25 (withdrawn): Use of the recombinant, non-human animal according to claim 24 for screening, testing, and validating compounds, agents, and modulators in the development of diagnostics and therapeutics to treat neurodegenerative diseases, in particular Alzheimer's disease.
- 26 (withdrawn): An assay for screening for a modulator of neurodegenerative diseases, in particular Alzheimer's disease, or related diseases or disorders of one or more substances selected from the group consisting of
- (i) a gene coding for hTARPP, and/or
 - (ii) a transcription product of the gene coding for hTARPP, and/or
 - (iii) a translation product of the gene coding for hTARPP, and/or
 - (iv) a fragment, or derivative, or variant of (i) to (iii),
- said method comprising:
- (a) contacting a cell with a test compound;

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- (b) measuring the activity and/or level of one or more substances recited in (i) to (iv);
- (c) measuring the activity and/or level of one or more substances recited in (i) to (iv) in a control cell not contacted with said test compound; and
- (d) comparing the levels and/or activities of the substance in the cells of step (b) and (c), wherein an alteration in the activity and/or level of substances in the contacted cells indicates that the test compound is a modulator of said diseases or disorders.

Claims 27-31 (cancelled).